SEQUENCE LISTING

<110> Sprecher, Cindy A. Kisiel, Walter Foster, Donald C.

<120> NOVEL HUMAN KUNITZ-TYPE INHIBITORS AND METHODS RELATING THERETO

<130> 93-14D4 <150> US 09/904,621 <151> 2001-07-13 <150> US 09/265,627 <151> 1999-03-09 <150> US 08/457,887 <151> 1995-06-01 <150> US 08/147,710 <151> 1993-11-05 <160> 15 <170> FastSEQ for Windows Version 4.0 <210> 1 <211> 979 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (39)...(746) <400> 1 ggacgcettg cccageggge egecegacee cetgeace atg gac eee get ege eee 56 Met Asp Pro Ala Arg Pro ctg ggg ctg tcg att ctg ctg ctt ttc ctg acg gag gct gca ctg ggc 104 Leu Gly Leu Ser Ile Leu Leu Phe Leu Thr Glu Ala Ala Leu Gly 10 15 gat gct gct cag gag cca aca gga aat aac gcg gag atc tgt ctc ctg 152 Asp Ala Ala Gln Glu Pro Thr Gly Asn Asn Ala Glu Ile Cys Leu Leu ccc cta gac tac gga ccc tgc cgg gcc cta ctt ctc cgt tac tac tac 200 Pro Leu Asp Tyr Gly Pro Cys Arg Ala Leu Leu Leu Arg Tyr Tyr Tyr 40 45 248 gac agg tac acg cag agc tgc cgc cag ttc ctg tac ggg ggc tgc gag Asp Arg Tyr Thr Gln Ser Cys Arg Gln Phe Leu Tyr Gly Gly Cys Glu 55 ggc aac gcc aac aat ttc tac acc tgg gag gct tgc gac gat gct tgc 296

Gly Asn Ala Asn Asn Phe Tyr Thr Trp Glu Ala Cys Asp Asp Ala Cys

75

80

tgg Trp	agg Arg	ata Ile	gaa Glu 90	aaa Lys	gtt Val	ccc Pro	aaa Lys	gtt Val 95	tgc Cys	cgg Arg	ctg Leu	caa Gln	gtg Val 100	agt Ser	gtg Val	344
gac Asp	gac Asp	cag Gln 105	tgt Cys	gag Glu	Gly ggg	tcc Ser	aca Thr 110	gaa Glu	aag Lys	tat Tyr	ttc Phe	ttt Phe 115	aat Asn	cta Leu	agt Ser	392
tcc Ser	atg Met 120	aca Thr	tgt Cys	gaa Glu	aaa Lys	ttc Phe 125	ttt Phe	tcc Ser	ggt Gly	Glà aaa	tgt Cys 130	cac His	cgg Arg	aac Asn	cgg Arg	440
att Ile 135	gag Glu	aac Asn	agg Arg	ttt Phe	cca Pro 140	gat Asp	gaa Glu	gct Ala	act Thr	tgt Cys 145	atg Met	ggc Gly	ttc Phe	tgc Cys	gca Ala 150	488
cca Pro	aag Lys	aaa Lys	att Ile	cca Pro 155	tca Ser	ttt Phe	tgc Cys	tac Tyr	agt Ser 160	cca Pro	aaa Lys	gat Asp	gag Glu	gga Gly 165	ctg Leu	536
tgc Cys	tct Ser	gcc Ala	aat Asn 170	gtg Val	act Thr	cgc Arg	tat Tyr	tat Tyr 175	ttt Phe	aat Asn	cca Pro	aga Arg	tac Tyr 180	aga Arg	acc Thr	584
tgt Cys	gat Asp	gct Ala 185	ttc Phe	acc Thr	tat Tyr	act Thr	ggc Gly 190	tgt Cys	gga Gly	GJÀ aaa	aat Asn	gac Asp 195	aat Asn	aac Asn	ttt Phe	632
gtt Val	agc Ser 200	agg Arg	gag Glu	gat Asp	tgc Cys	aaa Lys 205	cgt Arg	gca Ala	tgt Cys	gca Ala	aaa Lys 210	gct Ala	ttg Leu	aaa Lys	aag Lys	680
aaa Lys 215	aag Lys	aag Lys	atg Met	cca Pro	aag Lys 220	ctt Leu	cgc Arg	ttt Phe	gcc Ala	agt Ser 225	aga Arg	atc Ile	cgg Arg	aaa Lys	att Ile 230	728
	gg aag aag caa ttt taa acattettaa tatgteatet tgtttgtett rg Lys Lys Gln Phe * 235															776
atc ttt	tatggcttat ttgcctttat ggttgtatct gaagaataat atgacagcat gaggaaacaa atcattggtg atttattcac cagtttttat taatacaagt cactttttca aaaatttgga tttttttata tataactagc tgctattcaa atgtgagtct accattttta atttatggtt caactgtttg tgagactgaa ttc															896
<21 <21	<210> 2 <211> 235 <212> PRT <213> Homo sapiens															
	0> 2		λla	Ara	Pro	I.eu	Glv	Leu	Ser	·Tle	Leu	Leu	Leu	. Phe	Leu	
1				5					10					15	Asn	
			20					25					30		Leu	
		35	-				40					45			Phe	
	50					55					60				Glu	
65					70					75					80	
ATS	. Суя	ASP) ASP) Ala	cys	TIE	Arg	TTE	: GIU	r nys	val	. PIC	, nys	, val	. Cys	

```
85
                                     90
                                                         95
Arg Leu Gln Val Ser Val Asp Asp Gln Cys Glu Gly Ser Thr Glu Lys
            100
                                105
                                                     110
Tyr Phe Phe Asn Leu Ser Ser Met Thr Cys Glu Lys Phe Phe Ser Gly
                            120
                                                 125
Gly Cys His Arg Asn Arg Ile Glu Asn Arg Phe Pro Asp Glu Ala Thr
    130
                        135
                                             140
Cys Met Gly Phe Cys Ala Pro Lys Lys Ile Pro Ser Phe Cys Tyr Ser
                                                             160
                    150
                                         155
Pro Lys Asp Glu Gly Leu Cys Ser Ala Asn Val Thr Arg Tyr Tyr Phe
                165
                                     170
                                                         175
Asn Pro Arg Tyr Arg Thr Cys Asp Ala Phe Thr Tyr Thr Gly Cys Gly
            180
                                185
                                                     190
Gly Asn Asp Asn Asn Phe Val Ser Arg Glu Asp Cys Lys Arg Ala Cys
       195
                            200
                                                 205
Ala Lys Ala Leu Lys Lys Lys Lys Met Pro Lys Leu Arg Phe Ala
                        215
                                             220
Ser Arg Ile Arg Lys Ile Arg Lys Lys Gln Phe
                    230
                                         235
<210> 3
<211> 30
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide ZC4792
<400> 3
                                                                    30
attattacta ttacctccac agcctccata
<210> 4
<211> 30
<212> DNA
<213> Artificial Sequence
<220>
<223> Oligonucleotide ZC6281
<400> 4
                                                                    30
acagatetee gegttattte etgttggete
<210> 5
<211> 38
<212> DNA
<213> Artificial Sequence
<220>
<223> Clone M-2161
<400> 5
                                                                    38
gctgagagat tggagaagag agagatctgt ctcctgcc
<210> 6
<211> 34
<212> DNA
<213> Artificial Sequence
<220>
<223> M-2177
```

```
<400> 6
gaaacctcta gacttatatc ctccagcaag catc
                                                                    34
<210> 7
<211> 235
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> (77)...(235)
<400> 7
gaattccatt caagaatagt tcaaacaaga agattacaaa ctatcaattt catacacaat 60
ataaacgacc aaaaga atg aag gct gtt ttc ttg gtt ttg tcc ttg atc gga 112
                  Met Lys Ala Val Phe Leu Val Leu Ser Leu Ile Gly
                                    5
ttc tgc tgg gcc caa cca gtc act ggc gat gaa tca tct gtt gag att
                                                                    160
Phe Cys Trp Ala Gln Pro Val Thr Gly Asp Glu Ser Ser Val Glu Ile
         15
                              20
                                                  25
ccg gaa gag tct ctg atc atc gct gaa aac acc act ttg gct aac gtc
                                                                    208
Pro Glu Glu Ser Leu Ile Ile Ala Glu Asn Thr Thr Leu Ala Asn Val
     30
                         35
                                              40
gcc atg gct gag aga ttg gag aag aga
                                                                    235
Ala Met Ala Glu Arg Leu Glu Lys Arg
45
                     50
<210> 8
<211> 53
<212> PRT
<213> Homo sapiens
<400> 8
Met Lys Ala Val Phe Leu Val Leu Ser Leu Ile Gly Phe Cys Trp Ala
                                     10
                                                          15
Gln Pro Val Thr Gly Asp Glu Ser Ser Val Glu Ile Pro Glu Glu Ser
            20
                                 25
                                                     30
Leu Ile Ile Ala Glu Asn Thr Thr Leu Ala Asn Val Ala Met Ala Glu
                             40
                                                 45
Arg Leu Glu Lys Arg
   50
<210> 9
<211> 17
<212> DNA
<213> Artificial Sequence
<220>
<223> Nor-1478
<400> 9
gtaaaacgac ggccagt
                                                                    17
<210> 10
<211> 21
<212> DNA
<213> Artificial Sequence
```

```
<220>
<223> NOR-2523
<400> 10
                                                                   21
tctcttctcc aatctctcag c
<210> 11
<211> 35
<212> DNA
<213> Artificial Sequence
<220>
<223> M-2162
<400> 11
                                                                    35
cttttactct agacttactt tggtgcgcag aagcc
<210> 12
<211> 10
<212> PRT
<213> Artificial Sequence
<220>
<223> N-terminal
Asp Ala Ala Gln Glu Pro Thr Gly Asn Asn
<210> 13
<211> 8
<212> PRT
<213> Artificial Sequence
<220>
<223> N-terminal
<400> 13
Ala Gln Glu Pro Thr Gly Asn Asn
<210> 14
<211> 165
<212> DNA
<213> Artificial Sequence
<220>
<221> variation
<222> (0)...(0)
<223> Kunitz domain
<221> variation
<222> (1)...(3)
<223> Codon-1 Wherein the nucleotide triplet 1-3
      encodes any amino acid except cysteine.
<221> variation
<222> (4)...(6)
<223> Codon-2 Wherein the nucleotide triplet 4-6
```

encodes any amino acid except cysteine.

Cys Asp Asp Ala Cys Xaa Xaa

```
<221> variation
<222> (160)...(162)
<223> Codon-54 Wherein the nucleotide triplet 160-162
      encodes any amino acid except cysteine.
<221> variation
<222> (163)...(165)
<223> Codon-55 Wherein the nucleotide triplet 163-165
      encodes any amino acid except cysteine.
<400> 14
nnnnntgtc tcctgcccct agactacgga ccctgccggg ccctacttct ccgttactac 60
tacgacaggt acacgcagag ctgccgccag ttcctgtacg ggggctgcga gggcaacgcc 120
aacaatttct acacctggga ggcttgcgac gatgcttgcn nnnnn
<210> 15
<211> 55
<212> PRT
<213> Artificial Sequence
<220>
<223> Kunitz Domain
<221> VARIANT
<222> (1)...(2)
<223> aa1-2 Wherein each amino acid from position 1 to
      2 is individually any amino acid except cysteine.
<221> VARIANT
<222> (54)...(55)
<223> aa54-55 Wherein each amino acid from position 54
      to 55 is individually any amino acid except
      cysteine.
<400> 15
Xaa Xaa Cys Leu Pro Leu Asp Tyr Gly Pro Cys Arg Ala Leu Leu
                                    10
Leu Arg Tyr Tyr Tyr Asp Arg Tyr Thr Gln Ser Cys Arg Gln Phe Leu
            2.0
                                25
                                                    30
Tyr Gly Gly Cys Glu Gly Asn Ala Asn Asn Phe Tyr Thr Trp Glu Ala
```